

**AMENDMENTS TO THE CLAIMS:**

This listing of claims will replace all prior versions, and listings, of claims in the application:

**LISTING OF CLAIMS:**

Claim 1. (Currently Amended) A method of preparing a viral preparation predominantly containing adenoviruses, said method comprising:

- a) producing the viral preparation in an appropriate cell line;
- b) harvesting the viral preparation from the producer cell line and/or the culture supernatant;
- c) optionally, breaking the cells of the producer cell line;
- d) optionally clarifying;
- e) inactivating enveloped viruses in a the said viral preparation containing adenoviruses in which a sufficient quantity of a solvent comprising tri-n-butyl phosphate (TNBP) of between 0.1% and 0.6% (volume/volume) and TWEEN® 80 at a concentration of between 0.5% and 2% (volume/volume) are is introduced into the said viral preparation and the said solvent and TWEEN® 80 are is allowed to act at a temperature between +4°C to +37°C, at a pH of between 6.5 to 8.5 for a period which is sufficiently long to significantly reduce the quantity of enveloped viruses present in the said viral preparation, wherein said ~~method of~~ inactivation is capable of preserving at least 80% of the infectious activity of said adenoviruses; and
- f) optionally purifying.

Claims 2 – 9. (Canceled)

<sup>2</sup>  
Claim ~~10~~. (Currently Amended) The method of preparing a viral

preparation ~~inactivating enveloped viruses~~ according to Claim 1, wherein the solvent is allowed to act for a period of between 15 minutes and 24 hours.

<sup>3</sup>  
Claim ~~11~~. (Currently Amended) The method of preparing a viral

preparation ~~inactivating enveloped viruses~~ according to Claim 1, wherein the inactivating step method is carried out with stirring.

<sup>4</sup>  
Claim ~~12~~. (Currently Amended) The method of preparing a viral

preparation ~~inactivating enveloped viruses~~ according to Claim 1, wherein the inactivating step method is carried out under conductivity conditions of between 5 and 500 mS/cm.

Claims 13-17. (Canceled)

<sup>5</sup>  
Claim ~~18~~. (Currently Amended) The method of preparing a viral preparation according to ~~inactivating enveloped viruses of claim 1~~, wherein the quantity of TNBP introduced into the said viral preparation is ~~in the region of~~ about 0.3% (volume/volume).

Claims 19-21. (Canceled)

<sup>6</sup>  
Claim ~~22~~. (Currently Amended) The method of preparing a viral preparation according to inactivating enveloped viruses of claim 1, wherein the inactivating step is carried out at a the temperature of is between +15°C and +25°C.

<sup>7</sup>  
Claim ~~23~~. (Currently Amended) The method of preparing a viral preparation according to inactivating enveloped viruses of claim 1, wherein the inactivating step is carried out at pH is 8.5.

<sup>8</sup>  
Claim ~~24~~. (Currently Amended) The method of preparing a viral preparation according to inactivating enveloped viruses of claim <sup>2</sup>~~10~~, wherein the period of time is of between 1 hour and 5 hours.

<sup>9</sup>  
Claim ~~25~~. (Currently Amended) The method of preparing a viral preparation according to inactivating enveloped viruses of claim <sup>4</sup>~~12~~, wherein the conductivity conditions are between 10 and 100 mS/cm.

<sup>10</sup>  
Claim ~~26~~. (Currently Amended) The method of preparing a viral preparation according to inactivating enveloped viruses of claim 1, wherein TNBP at a final concentration of between 0.1% and 0.6% (volume/volume) and Tween<sup>®</sup>-80 at a final concentration of between 0.5% and 2% (volume/volume) are introduced into said viral preparation, said TNBP and said Tween<sup>®</sup>-80 are allowed to act at room temperature at a pH of 8.5 for a period of time between 1 hour and 5 hours, wherein at least 80% of the infectious activity of said adenoviruses is preserved.

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Claim ~~27~~. (Currently Amended) The method of preparing a viral preparation according to inactivating enveloped viruses of claim 1, wherein said adenovirus is recombinant.

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Claim ~~28~~. (Currently Amended) The method of preparing a viral preparation according to inactivating enveloped viruses of claim 1, wherein said adenovirus is replication-defective.

Claims 29-31. (Canceled)

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Claim ~~32~~. (New) The method of preparing a viral preparation according to claim 1, wherein in the inactivating step (e), the final concentration of TNBP is 0.6% and the final concentration of TWEEN® 80 is 2%.

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Claim ~~33~~. (New) The method of preparing a viral preparation according to claim 1, wherein in the inactivating step (e), the final concentration of TNBP is 0.3% and the final concentration of TWEEN® 80 is 1%.

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Claim ~~34~~. (New) The method of preparing a viral preparation according to claim 1, wherein the producing of the viral preparation is from a cell line having undergone an infection carried out at a multiplicity of infection (MOI) of about 1 to 10.

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Claim ~~35~~. (New) The method of preparing a viral preparation according to claim 1, wherein the viral preparation is harvested from the producing cell line and from the culture supernatant.

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Claim ~~36~~<sup>16</sup>. (New) The method of preparing a viral preparation according to claim 1 or ~~35~~<sup>16</sup>, wherein the viral preparation is harvested at 48h or 72h post-infection.

~~36~~<sup>18</sup>

Claim ~~37~~<sup>18</sup>. (New) The method of preparing a viral preparation according to claim 1, wherein the breaking of the cells of the producing cell line is carried out by a technique selected from a group consisting of freeze-thaw cycles, enzymatic lysis, chemical means, mechanical means, and any combination thereof.

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Claim ~~38~~<sup>18</sup>. (New) The method of preparing a viral preparation according to claim ~~37~~<sup>18</sup>, wherein the mechanical means is selected from the group consisting of ultrasound, attrition, pressure and shear forces, microfluids, the mechanical action of two rollers generating hydraulic and mechanical shear forces, and any combination thereof.

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Claim ~~39~~<sup>20</sup>. (New) The method of preparing a viral preparation according to claim 1, wherein the clarification step comprises successive filtrations carried out on filters of decreasing porosity.

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Claim ~~40~~<sup>20</sup>. (New) The method of preparing a viral preparation according to claim ~~39~~<sup>20</sup>, wherein the successive filtrations are carried out on a filter having a porosity of 8  $\mu\text{m}$ , then on a filter having a porosity of 5  $\mu\text{m}$ , then on a filter having a porosity of between 3  $\mu\text{m}$  and 0.8  $\mu\text{m}$ , and then on a filter having a porosity of between 0.8  $\mu\text{m}$  and 0.65  $\mu\text{m}$ .

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Claim ~~41~~. (New) The method of preparing a viral preparation according to claim 1, wherein the clarification step comprises tangential microfiltration.

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Claim ~~42~~. (New) The method of preparing a viral preparation according to claim 1, wherein the purification step comprises an ion-exchange chromatographic step.

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Claim ~~43~~. (New) The method of preparing a viral preparation according to claim ~~42~~,<sup>23</sup> wherein the purification step comprises an ion-exchange chromatography and a gel filtration chromatography.

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Claim ~~44~~. (New) The method of preparing a viral preparation according to claim ~~42~~,<sup>23</sup> wherein the ion-exchange chromatography is carried out on a support functionalized with quaternary amines.

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Claim ~~45~~. (New) The method of preparing a viral preparation according to claim ~~43~~,<sup>24</sup> wherein the gel filtration chromatography is carried out on a support selected from the group consisting of supports comprising allyl dextran-methylene bisacrylamide matrices, ethylene glycol-methacrylate matrices, N-acrylamine hydroxypropanediol matrices and agarose matrices.

<sup>27</sup>  
Claim ~~46~~. (New) The method of preparing a viral preparation according to claim 1, wherein said method further comprises a step of degrading nucleic acids.

<sup>28</sup>  
Claim ~~47~~. (New) The method of preparing a viral preparation according to claim ~~46~~,<sup>27</sup> wherein the step of degrading nucleic acids comprises treatment with benzonase.

<sup>29</sup>  
Claim ~~48~~. (New) The method of preparing a viral preparation according to claim ~~47~~,<sup>28</sup> wherein the benzonase treatment and the inactivating step (e) are carried out simultaneously after breaking (c) and clarification (d) steps.